

ISIS-4718

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

*Pre Amended  
2A*

In re patent application of:

Phillip D. Cook and Muthiah Manoharan

Serial No.: Not yet assigned

Group No.: Not yet assigned

Filed: Herewith

Examiner: Not yet assigned

For: Phosphorothioate Oligonucleotides Having Modified  
Internucleoside Linkages

Assistant Commissioner for  
Patents  
Washington, DC 20231

PRELIMINARY AMENDMENT

Prior to examination of the above referenced application, Applicants respectfully request entry of the following amendment, without prejudice:

In the claims:

Please cancel claims 1 and 16.

Please amend claims 2-15 and 17-19 as indicated below:

*2*  
2. The method of claim ~~10~~ wherein  $L_1$  is a Sp phosphorothioate internucleoside linkage and  $L_2$  is a racemic phosphorothioate internucleoside linkage.

3. The method of claim ~~10~~ wherein  $L_1$  and  $L_2$  both are Sp phosphorothioate internucleoside linkages.

4. The method of claim ~~10~~ wherein  $L_1$  is a Rp phosphorothioate